

## CLAIMS

1        1. A method of achieving a therapeutic effect comprising:  
2                providing a particle containing a therapeutic substance to an anatomical structure  
3                comprising a lumen such that said particle embolizes within said lumen for a transitory  
4                period of less than one week;  
5                wherein said therapeutic substance is released from said particle, causing said  
6                therapeutic effect.

1        2. The method of Claim 1, wherein said anatomical structure has a first region and a  
2        second region branching from said first region, said second region being located downstream  
3        from said first region, and wherein said providing a particle to an anatomical structure comprises  
4        the acts of:  
5                causing an occlusion in said first region at a position downstream of the location  
6                at which said second region branches from said first region; and  
7                introducing said particle into said first region upstream of the location at which  
8                said second region branches from said first region.

1        3. The method of Claim 1, wherein said anatomical structure has a first region and a  
2        second region branching from said first region, said second region being located downstream  
3        from said first region, and wherein said providing a particle to an anatomical structure comprises  
4        the acts of:  
5                occluding said first region at positions both upstream and downstream of the  
6                location at which said second region branches from said first region; and  
7                introducing said particle into said first region between the upstream and  
8                downstream occlusions.

1        4. The method of Claim 1, wherein said lumen contains an occlusion therein and  
2        wherein said providing a particle to an anatomical structure comprises the act of:  
3                introducing said particle into said lumen upstream of said occlusion.

1        5.     The method of Claim 4, wherein said therapeutic substance is an angiogenic  
2     substance and wherein said therapeutic effect is collateral growth upstream of said occlusion.

1        6.     The method of Claim 1, wherein said particle reduces in size as said therapeutic  
2     substance is released therefrom.

1        7.     A method according to Claim 1 wherein the method of providing a particle further  
2     comprises:

3            preparing a solution;  
4            emulsifying the solution to form an emulsion; and  
5            filtering particles from the emulsion.

1        8.     A method according to Claim 7 wherein:

2            preparing the solution further includes:  
3              dissolving lecithin and dexamethasone in methylene chloride;  
4              emulsifying the solution further includes:  
5                stirring the solution with a perfluorotributylamine / pluronic F-68 and water  
6                solution with heparin; and  
7              filtering particles further includes:  
8                warming the emulsion to drive off the methylene chloride to form microparticles;  
9                collecting the microparticles by filtration;  
10              washing the collected microparticles in cold water;  
11              drying the microparticles in a vacuum; and  
12              separating the microparticles by size using cyclone or tangential flow filtration.

1        9.     A method according to Claim 7 wherein:

2            preparing the solution further includes:  
3              dissolving Basic Fibroblast Growth Factor, heparin, and lactose in phosphate  
4              buffer with Human serum albumin;  
5              emulsifying the solution further includes:  
6                mixing the solution methylene chloride containing polylactide-co-glycolide to  
7                form a single emulsion;  
8                stirring the first emulsion with a perfluorotributylamine / pluronic F-68 and water  
9                solution to form a double emulsion; and

10 filtering particles further includes:

11 warming the emulsion to drive off the methylene chloride to form microparticles;  
12 collecting the microparticles by filtration;  
13 washing the collected microparticles in water;  
14 mixing the washed microparticles with an aqueous solution of mannose in  
15 potassium phosphate buffer;  
16 freeze-drying the microparticles in a vacuum; and  
17 separating the microparticles by size using cyclone or tangential flow filtration.

1 10. A method according to Claim 7 wherein:

2 preparing the solution further includes:

3 stirring plasmid DNA and heparin in a solution of dextran and mannose in water;  
4 emulsifying the solution further includes:

5 emulsifying the solution in cyclo-octane with SPAN 80; and

6 filtering particles further includes:

7 filling the emulsion into lyophilization vials to form microparticles;

8 freeze-drying the microparticles; and

9 separating the microparticles by size using cyclone or tangential flow filtration.

1 11. A method of achieving a therapeutic effect comprising:

2 providing a particle to an anatomical structure having a lumen such that said  
3 particle embolizes within said lumen for a transitory period;

4 wherein said transitory period of embolization causes a brief period of  
5 reduced blood flow through said lumen that induces a therapeutic bodily response.

1 12. The method of Claim 11, wherein said act of providing a particle to said

2 anatomical structure comprises the act of delivering pulses of said particles to said anatomical  
3 structure.

1 13. The method of Claim 12, wherein the act of delivering pulses of said particles

2 causes a series of said brief periods of reduced blood flow;

3 wherein said therapeutic bodily response induced by said series of brief periods of  
4 reduced blood flow is collateral growth.

1        14. A composition for achieving a therapeutic effect in an anatomical structure  
2 comprising a lumen, the composition comprising:

3              a particle suitable for introduction into an anatomical structure, said particle  
4 containing a therapeutic substance and being capable of reducing in size;

5              wherein said particle is capable of embolizing within said lumen for a  
6 transitory period of less than one week; and

7              wherein said therapeutic substance is released from said particle for the treatment  
8 of a patient.

,1        15. The composition of Claim 14, wherein said therapeutic substance is selected from  
2 a group of antineoplastic, antiplatelet, anticoagulant, fibrinolytic, antimitotic, thrombin inhibitor,  
3 antiinflammatory, antiproliferative, antioxidant, antiangiogenic, angiogenic, arteriogenic,  
4 antiallergic substances, and mixtures thereof.

1        16. The composition of Claim 14, wherein said particle is made of a mixture of at  
2 least two different substances.

1        17. The composition of Claim 16, wherein each of said substances reduces in size in  
2 said lumen at a different rate.

1        18. The composition of Claim 14, wherein said particle is made of a first substance  
2 and a second substance, said second substance covering at least a portion of said first substance,  
3 wherein each of said substances reduce in size in said lumen at a different rate.

1        19. The composition of Claim 14, wherein said particle reduces in size as said  
2 therapeutic substance is released therefrom.

1        20. A composition for achieving a therapeutic effect in an anatomical structure  
2 comprising a lumen, said composition comprising:  
3              a particle suitable for introduction into an anatomical structure, said particle being  
4 capable of reducing in size;

5                   wherein said particle is capable of embolizing within said lumen for a  
6                   transitory period, causing a brief period of reduced blood flow which induces a  
7                   therapeutic bodily response.

1                 21. A method of achieving a therapeutic effect within an anatomical structure having  
2                 a first region and a second region, said second region being located downstream of said first  
3                 region and having a smaller cross-sectional diameter than said first region, the method  
4                 comprising the acts of:

- 5                 (a) providing a particle having a first size in which said particle is not capable of  
6                 passing from said first region into said second region, said particle being capable of  
7                 reducing in size; and  
8                 (b) delivering said particle having said first size to said first region of said  
9                 anatomical structure;

10                 wherein said particle subsequently reduces from said first size to a smaller second  
11                 size as said particle travels through said anatomical structure, allowing said particle to  
12                 pass into said second region; and

13                 wherein a therapeutic effect is achieved.

1                 22. The method of Claim 21, wherein said particle includes a therapeutic substance;  
2                 wherein said therapeutic substance is released from said particle; and  
3                 wherein said therapeutic effect results from said therapeutic substance.

1                 23. The method of Claim 21, wherein during said act of traveling through said  
2                 anatomical structure and prior to said act of reducing to said second size, said particle reaches a  
3                 diameter of said anatomical structure through which said particle cannot pass and at which said  
4                 particle is constrained for a transitory period until said particle reduces to said second size.

1                 24. The method of Claim 23, wherein said particle includes a therapeutic substance  
2                 and wherein said transitory period is less than one week;  
3                 wherein said therapeutic substance is released from said particle; and  
4                 wherein said therapeutic effect results from said therapeutic substance.

1        26.     The method of Claim 24, wherein said particle reduces in size as said therapeutic  
2 substance is released therefrom.

1        27.     The method of Claim 21, wherein during said act of traveling through said  
2 anatomical structure, said particle becomes transiently lodged in a plurality of locations  
3 throughout said anatomical structure as said particle reduces in size over a period of days,  
4 providing said therapeutic effect over a length of said anatomical structure.

1        28.     The method of Claim 21, wherein said anatomical structure is within a  
2 mammalian cardiovascular system,

3                wherein a brief period of reduced blood flow is caused during said transitory  
4 period; and

5                wherein said therapeutic effect is a therapeutic bodily response induced by said  
6 brief period of reduced blood flow.

1        29.     The method of Claim 21, wherein said anatomical structure comprises a single  
2 lumen containing said first region and said second region.

1        30.     The method of Claim 21, wherein said anatomical structure comprises a lumen  
2 network including a plurality of lumens.

2        31.     The method of Claim 21, wherein said anatomical structure additionally includes  
3 a third region, said third region being located downstream of said second region and having a  
4 smaller cross-sectional diameter than said second region;

5                wherein said particle is capable of reducing from said second size to a smaller  
6 third size, allowing said particle to pass from said second region into said third region.